

GenCore version 5.1.6  
Copyright (c) 1993 - 2003 Compugen Ltd.

OM protein - protein search, using sw model

Run on: June 25, 2003, 14:20:41 ; Search time 33.3 Seconds

(without alignments)  
444.169 Million cell updates/sec

Title: us-09-622-613b-21

Sequence: 1 MONMATEFOOKHINTEPLICN.....ICVKCENQYVPHFGIGRCP 111

Scoring table:

BLOSUM62  
Gapop 10.0 , Gapext 0.5

Searched: 908470 seqs, 133250620 residues

Total number of hits satisfying chosen parameters: 908470

Minimum DB seq length: 10

Maximum DB seq length: 2000000000

Post-processing: Minimum Match 0%

Maximum Match 100%  
Listing first 45 summaries

Database :

1: /SID52/gcgdata/geneseq/geneseq-emb1/AA1980.DAT:\*  
2: /SID52/gcgdata/geneseq/geneseq-emb1/AA1981.DAT:\*  
3: /SID52/gcgdata/geneseq/geneseq-emb1/AA1982.DAT:\*  
4: /SID52/gcgdata/geneseq/geneseq-emb1/AA1983.DAT:\*  
5: /SID52/gcgdata/geneseq/geneseq-emb1/AA1984.DAT:\*  
6: /SID52/gcgdata/geneseq/geneseq-emb1/AA1985.DAT:\*  
7: /SID52/gcgdata/geneseq/geneseq-emb1/AA1986.DAT:\*  
8: /SID52/gcgdata/geneseq/geneseq-emb1/AA1987.DAT:\*  
9: /SID52/gcgdata/geneseq/geneseq-emb1/AA1988.DAT:\*  
10: /SID52/gcgdata/geneseq/geneseq-emb1/AA1989.DAT:\*  
11: /SID52/gcgdata/geneseq/geneseq-emb1/AA1990.DAT:\*  
12: /SID52/gcgdata/geneseq/geneseq-emb1/AA1991.DAT:\*  
13: /SID52/gcgdata/geneseq/geneseq-emb1/AA1992.DAT:\*  
14: /SID52/gcgdata/geneseq/geneseq-emb1/AA1993.DAT:\*  
15: /SID52/gcgdata/geneseq/geneseq-emb1/AA1994.DAT:\*  
16: /SID52/gcgdata/geneseq/geneseq-emb1/AA1995.DAT:\*  
17: /SID52/gcgdata/geneseq/geneseq-emb1/AA1996.DAT:\*  
18: /SID52/gcgdata/geneseq/geneseq-emb1/AA1997.DAT:\*  
19: /SID52/gcgdata/geneseq/geneseq-emb1/AA1998.DAT:\*  
20: /SID52/gcgdata/geneseq/geneseq-emb1/AA1999.DAT:\*  
21: /SID52/gcgdata/geneseq/geneseq-emb1/AA2000.DAT:\*  
22: /SID52/gcgdata/geneseq/geneseq-emb1/AA2001.DAT:\*  
23: /SID52/gcgdata/geneseq/geneseq-emb1/AA2002.DAT:\*

Pred. No. is the number of results predicted by chance to have a score greater than or equal to the score of the result being printed, and is derived by analysis of the total score distribution.

# SUMMARIES

Result No.	Score	Query Match	Length	DB ID	Description
1	605	100.0	111	20	AAV28876
2	601	99.3	111	20	AAV28873
3	600	99.2	110	20	AAV28874
4	596	98.5	110	20	AAV28872
5	596	98.5	111	20	AAV28878
6	591	97.7	110	20	AAV28877
7	582.5	96.3	111	20	AAV33321
8	284.5	47.0	105	20	AAV28869
9	282.5	46.7	105	20	AAV28867
10	280.5	46.4	104	18	AAW06544

11	279.5	46.2	104	20	AAV28866	Recombinant RAPRI
12	278.5	46.0	105	20	AAV39400	Recombinant frog O
13	277.5	45.9	104	20	AAV28865	Rana pipiens liver
14	277.5	45.9	105	20	AAV28871	Recombinant Met(-1
15	277.5	45.9	127	20	AAV28879	Rana pipiens Clone
16	275.5	45.5	105	18	AAW5123	R. pipiens recombi
17	275.5	45.5	355	18	AAW5125	R. pipiens recombi
18	275.5	45.5	358	18	AAW5130	R. pipiens recombi
19	273.5	45.2	104	18	AAW30301	Recombinant one pr
20	273.5	45.2	104	22	AAW31666	Amino acid sequenc
21	273.5	45.2	104	22	AAW31667	Amino acid sequenc
22	273.5	45.2	112	18	AAW5118	R. pipiens recombi
23	273.5	45.2	251	18	AAW5134	R. pipiens recombi
24	273.5	45.2	254	18	AAW5135	R. pipiens recombi
25	273.5	45.2	355	18	AAW5129	R. pipiens recombi
26	273.5	45.2	355	18	AAW5133	R. pipiens recombi
27	273.5	45.2	366	18	AAW5132	R. pipiens recombi
28	273.5	45.2	379	18	AAW5136	R. pipiens recombi
29	272.5	45.0	104	18	AAW50302	Recombinant one pr
30	272.5	45.0	104	20	AAV28870	Recombinant RAPRI
31	270.5	44.7	104	12	AAW12343	protein with activ
32	270.5	44.7	104	15	AAW47303	ONCONASE (pharmac
33	270.5	44.7	104	17	AAW0736	Protein derived fr
34	270.5	44.7	104	18	AAW06543	Antitumor protein
35	270.5	44.7	104	18	AAW14065	Onconase (RTM) pro
36	270.5	44.7	104	20	AAV33322	Frog onconase prot
37	270.5	44.7	104	20	AAW88233	Rana pipiens RNase
38	268.5	44.4	105	18	AAW5116	R. pipiens recombi
39	268.5	44.4	106	18	AAW5122	R. pipiens recombi
40	268.5	44.4	107	18	AAW5117	R. pipiens recombi
41	267.5	44.2	105	18	AAW5115	R. pipiens recombi
42	264.5	43.7	358	18	AAW5127	R. pipiens recombi
43	264.5	43.7	365	18	AAW5131	R. pipiens recombi
44	263.5	43.6	104	18	AAW18224	Antitumor generic
45	246.5	40.7	107	18	AAW5120	R. pipiens recombi

# ALIGNMENTS

RESULT 1	
AAV28876	
ID	AAV28876 standard; Protein: 111 AA.
XX	
AC	AAV28876:
XX	
DT	25-JAN-2000 (first entry)
XX	
DE	Recombinant Met(-1) RacOR1 Met22Leu Met57Leu-(His)6 protein.
XX	
KW	Met(-1) Rana catesbeiana ribonuclease Met22Leu Met57Leu-(His)6. RacOR1
KW	recombinant; CD22: covalently bound; LL2 antibody; ligand binding molecu
KW	cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;
KW	signal peptide; recombinant ribonuclease; cytotoxic fusion protein;
KW	cancer; bullfrog; RNase; autoimmune disease.
XX	
OS	Rana catesbeiana.
OS	Synthetic.
XX	
FT	Key
FT	Misc-difference 1
FT	/note= "Location/Qualifiers
FT	/note= "(His)6 histidine tag attached to N-terminal Met"
FT	Misc-difference 1
FT	/note= "Met not found in wild type RacOR1"
FT	Misc-difference 23
FT	/note= "wild type Met replaced with Leu"
FT	Misc-difference 58
FT	/note= "wild type Met replaced with Leu"
XX	
PN	W09950398-A2.
XX	
PD	07-OCT-1999.
XX	

XX	27-MAR-1998;	98US-0079751.
XX	(USSH ) US DEPT HEALTH & HUMAN SERVICES.	
PA	Newton DL, Rybak SM;	
PI	WPI: 1999-610847/52.	
DR	N-PDSB; AAZ08131.	
DR		
XX		
PT	New recombinant ribonucleases, used for killing target cells, e.g. for treating cancers, viral infections or autoimmune diseases	-
PT		
XX	Claim 22; Page 63; 71pp; English.	
XX		
CC	The present sequence is a recombinant Rana catesbeiana oocyte	
CC	ribonuclease (RACOR1) protein with Met at position 1. Carboxy terminal	
CC	end of recombinant RACOR1 has a covalently bound ligand binding moiety,	
CC	which can be a LL2 antibody directed against CD22 on cancerous B cells or	
CC	human chorionic gonadotropin (hCG) effective against Kaposi's sarcoma	
CC	cells. Recombinant ribonucleases can be expressed in bacteria without an	
CC	N-terminal methionine due to the presence of a signal peptide that is	
CC	cleaved by bacteria. The soluble expression of ribonuclease allows the	
CC	proteins to be fused in-frame with ligand binding moieties to form	
CC	cytotoxic fusion proteins. They can be used for treatment of cancer and	
CC	autoimmune diseases.	
SQ	Sequence 111 AA;	
	Query Match 99.3%; Score 601; DB 20; Length 111;	
	Best Local Similarity 98.2%; Pred. No. 2e-60; Mismatches 0; Gaps 0;	
	Matches 109; Conservative 2; Indels 0;	
OY	1 MNMNAFQQKHIINFTPICNTILDNNIYIVGGCKRVNTEFISSATVKAICTGVINLV 60	
DY	1 MNMNAFQQKHIINFTPICNTIMNNIYIVGGCKRVNTEFISSATVKAICTGVINNV 60	
OY	61 LSTRPFLNLCTRTSTIRPPCPYSRRFTNTICYKCENQYVHRAGIGRCP 111	
DY	61 LSTRPFLNLCTRTSTIRPPCPYSRRFTNTICYKCENQYVHRAGIGRCP 111	
RESULT 3		
AAAY28874		
ID	AAV28874 standard; Protein: 110 AA.	
XX	AAV28874;	
AC		
XX		
DT	25-JAN-2000 (first entry)	
XX		
DE	Recombinant RACOR1 Met22Leu Met57Leu amino acid sequence.	
KX	Recombinant Rana catesbeiana oocyte ribonuclease; covalently bound;	
KW	RACOR1 Met22Leu Met57Leu; LL2 antibody; ligand binding moiety; CD22;	
KW	cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;	
KW	signal peptide; recombinant ribonuclease; cytotoxic fusion protein;	
KW	cancer; bullfrog; RNase; autoimmune disease.	
OS	Rana catesbeiana.	
OS	Synthetic.	
XX		
XX	Key Location/Qualifiers	
FT	Misc-difference 22	
FT	/note= "Wild type Met replaced with Leu"	
FT	Misc-difference 57	
FT	/note= "Wild type Met replaced with Leu"	
XX	WO950398-A2.	
PN		
PD	07-OCT-1999.	
XX		
XX	26-MAR-1999; 99WO-US0641.	
PE		
XX	27-MAR-1998; 98US-0079751.	

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PA Newlon DL, Rybak SM;  
 PI  
 XX  
 DR WPI: 1999-610847/52.  
 N-PSDB: AA208132.  
 XX  
 PT New recombinant ribonucleases, used for killing target cells, e.g. for  
 treating cancers, viral infections or autoimmune diseases -  
 XX  
 PS Claim 22: Page 64; 71pp; English.  
 CC The present sequence is a recombinant Rana catesbeiana oocyte  
 CC ribonuclease (RacOR1) protein with Met22Leu Met57Leu. Carboxy terminal  
 CC end of recombinant RacOR1 has a covalently bound ligand binding moiety,  
 CC which can be a IL2 antibody directed against CD22 on cancerous B cells  
 CC or human chorionic gonadotropin (hCG) effective against Kaposi's sarcoma  
 CC cells. Recombinant ribonucleases can be expressed in bacteria without an  
 CC N-terminal methionine due to the presence of a signal peptide that is  
 CC cleaved by bacteria. The soluble expression of ribonuclease allows the  
 CC proteins to be fused in-frame with ligand binding moieties to form  
 CC cytotoxic fusion proteins. They can be used for treatment of cancer and  
 CC autoimmune diseases.  
 CC  
 XX Sequence 110 AA:  
 S0  
 Query Match 99.2%; Score 600; DB 20; Length 110;  
 Best Local Similarity 100.0%; Pred. No. 2.5e-60;  
 Matches 110; Conservative 0; Mismatches 0; Indels 0; Gaps 0;  
 OY 2 QNNATFOQKHIIIMPICNTILDNNIYVGGCKRVNTFISSATVKAICTGVINLNL 61  
 Db 1 QNNATFOQKHIIIMPICNTILDNNIYVGGCKRVNTFISSATVKAICTGVINLNL 60  
 OY 62 STTRFQJLNTCTRTSITPRPCPYSSRTETNNICVCKENQYVHFAGIGRCP 111  
 Db 61 STTRFQJLNTCTRTSITPRPCPYSSRTETNNICVCKENQYVHFAGIGRCP 110  
 OY  
 Db  
 RESULT 4  
 AAY28872  
 ID AAY28872 standard; Protein: 110 AA.  
 XX  
 AC AAY28872;  
 XX  
 DT 25-JAN-2000 (first entry)  
 DE Rana catesbeiana oocyte ribonuclease (RacOR1) amino acid sequence.  
 XX  
 KM Rana catesbeiana oocyte ribonuclease; RacOR1: covalently bound; CD22;  
 KM IL2 antibody; ligand binding moiety; cancerous B cell; Kaposi's Sarcoma;  
 KM human chorionic gonadotropin; hCG; recombinant ribonuclease; bullfrog;  
 KM signal peptide; cytotoxic fusion protein; cancer; autoimmune disease;  
 KM  
 XX  
 OS Rana catesbeiana.  
 OS Synthetic.  
 OS  
 PN W09950398-A2.  
 XX  
 PD 07-OCT-1999.  
 XX  
 PF 26-MAR-1999; 99MO-US06641.  
 XX  
 PR 27-MAR-1998; 98US-0079751.  
 XX  
 PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PI Newlon DL, Rybak SM;  
 XX  
 DR WPI: 1999-610847/52.  
 N-PSDB: AA208130.

XX New recombinant ribonucleases, used for killing target cells, e.g. for  
 PT treating cancers, viral infections or autoimmune diseases -  
 XX  
 PS Claim 22: Page 62; 71pp; English.  
 CC The present sequence is a Rana catesbeiana oocyte ribonuclease (RacOR1)  
 CC protein encoded by a cDNA modified for expression in E. coli. Carboxy  
 CC terminal end of RacOR1 has a covalently bound ligand binding moiety,  
 CC which can be a IL2 antibody directed against CD22 on cancerous B cells  
 CC or human chorionic gonadotropin (hCG) effective against Kaposi's  
 CC Sarcoma cells. Recombinant ribonucleases can be expressed in bacteria  
 CC without an N-terminal methionine due to the presence of a signal peptide  
 CC that is cleaved by bacteria. The soluble expression of ribonuclease  
 CC allows the proteins to be fused in-frame with ligand binding moieties to  
 CC form cytotoxic fusion proteins. They can be used for treatment of cancer  
 CC and autoimmune diseases.  
 CC  
 XX Sequence 110 AA:  
 S0  
 Query Match 98.5%; Score 596; DB 20; Length 110;  
 Best Local Similarity 98.2%; Pred. No. 7.2e-60;  
 Matches 108; Conservative 2; Mismatches 0; Indels 0; Gaps 0;  
 OY 2 QNNATFOQKHIIIMPICNTILDNNIYVGGCKRVNTFISSATVKAICTGVINLNL 61  
 Db 1 QNNATFOQKHIIIMPICNTILDNNIYVGGCKRVNTFISSATVKAICTGVINLNL 60  
 OY 62 STTRFQJLNTCTRTSITPRPCPYSSRTETNNICVCKENQYVHFAGIGRCP 111  
 Db 61 STTRFQJLNTCTRTSITPRPCPYSSRTETNNICVCKENQYVHFAGIGRCP 110  
 OY  
 Db  
 RESULT 5  
 AAY28878  
 ID AAY28878 standard; Protein: 111 AA.  
 XX  
 AC AAY28878;  
 XX  
 DT 25-JAN-2000 (first entry)  
 DE Recombinant Met(-1) RacOR1 Gln1Ser amino acid sequence.  
 XX  
 KM Recombinant Met(-1) Rana catesbeiana oocyte ribonuclease Gln1Ser; RacOR1;  
 KM covalently bound; IL2 antibody; ligand binding moiety; cancerous B cell;  
 KM Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;  
 KM recombinant ribonuclease; cytotoxic fusion protein; cancer; bullfrog;  
 KM CD22; RNase; autoimmune disease.  
 KM  
 XX  
 OS Rana catesbeiana.  
 OS Synthetic.  
 OS  
 PN W09950398-A2.  
 XX  
 PD 07-OCT-1999.  
 XX  
 PF 26-MAR-1999; 99MO-US06641.  
 XX  
 PR 27-MAR-1998; 98US-0079751.  
 XX  
 PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 PI Newlon DL, Rybak SM;  
 XX  
 DR WPI: 1999-610847/52.  
 N-PSDB: AA208135.

PT	New recombinant ribonucleases, used for killing target cells, e.g. for treating cancers, viral infections or autoimmune diseases
XX	
XX	
XX	Claim 22; Page 68; 71pp; English.
CC	The present sequence is a recombinant Rana catesbeiana ribonuclease (RACOR1) protein with Met at position 1 and Gln2Ser. Carboxy terminal end of recombinant RACOR1 has a covalently bound ligand binding moiety, which can be a IL2 antibody directed against CD22 on cancerous B cells or human chorionic gonadotrophin (hCG) effective against Kaposi's sarcoma cells. CC Recombinant ribonucleases can be expressed in bacteria without an N-terminal methionine due to the presence of a signal peptide that is cleaved by bacteria. The soluble expression of ribonuclease allows the proteins to be fused in-frame with ligand binding moieties to form cytotoxic fusion proteins. They can be used for treatment of cancer and autoimmune diseases.
SO	Sequence 111 AA;
QY	Query Match 98.5%; Score 596; DB 20; Length 111; Best Local Similarity 97.3%; Pred. NO. 7.3e-60; Matches 108; Conservative 2; Mismatches 1; Indels 0; Gaps 0
Db	1 MONNATPQQKHIINPIICNTILDNNIIVGSGCKRVMTFTIISATYKAICTGYINLV 60   1 MSNNATPEQKHIIINPIICNTIMDNIIVGGCKRVMTFTIISATYKAICTGVINNV 60
OY	61 LSTRFOLNCTRTSITPRPCYSRTETNYICVCKENQYPVHFGIGRCP 111   61 LSTRFOLNCTRTSITPRPCYSRTETNYICVCKENQYPVHFGIGRCP 111
Db	
RESULT 6	
AAY28877	
ID	AAY28877 standard; Protein; 110 AA.
XX	
XX	AAY28877;
XX	
DT	25-JAN-2000 (first entry)
XX	
DE	Recombinant RACOR1 Gln1ser amino acid sequence.
KW	Recombinant Rana catesbeiana oocyte ribonuclease; RACOR1 Gln1ser; CD22; covalently bound; IL2 antibody; ligand binding moiety; cancerous B cell; bullfrog; Kaposi's sarcoma; human chorionic gonadotrophin; hCG; RNase; signal peptide; recombinant ribonuclease; cytotoxic fusion protein;
KM	cancer; autoimmune disease.
RW	
KW	Rana catesbeiana.
OS	Synthetic.
EH	Key Location/Qualifiers
FT	Misc-difference 1 /note= "Wild type Gln replaced with Ser"
PN	MO9950398-A2.
PD	07-OCT-1999.
PF	26-MAR-1999; 99WO-US06641.
PR	27-MAR-1998; 98US-0079751.
PA	(USSH ) US DEPT HEALTH & HUMAN SERVICES.
P1	Newton DL, Rybak SM;
WI	WPI: 1999-610847/52.
DR	N-PSDB; AAZ08134.
XX	
XX	New recombinant ribonucleases, used for killing target cells, e.g. for treating cancers, viral infections or autoimmune diseases
XX	

PS	Claim 22, Page 67; 71pp: English.
XX	
CC	The present sequence is a recombinant Rana catesbeiana oocyte
CC	ribonuclease (RacOR) protein with glutiser. Carboxy terminal end of
CC	recombinant RacOR1 has a covalently bound ligand binding moiety, which
CC	can be a IL2 antibody directed against CD22 on cancerous B cells or
CC	human chorionic gonadotropin (hCG) effective against Kaposi's sarcoma
CC	cells. Recombinant ribonucleases can be expressed in bacteria without an
CC	N-terminal methionine due to the presence of a signal peptide that is
CC	cleaved by bacteria. The soluble expression of a ribonuclease allows the
CC	proteins to be fused in-frame with ligand binding moieties to form
CC	cytotoxic fusion proteins. They can be used for treatment of cancer and
CC	autoimmune diseases.
XX	
SO	Sequence 110 AA:
	Query Match 97.7%; Score 591; DB 20; Length 110:
	Best Local Similarity 98.2%; Pred. No. 2.7e-59;
	Matches 107; Conservative 2; Mismatches 0; Indels 0; Gaps 0;
OY	3 NWATPQOKHIIINPTICMTILDNNIYIVGGCKRVNFTFISSATPYKAICTGVINLVLS 62
DB	2 NWATPQOKHIIINPTICMTINDNNIYIVGGCKRVNFTFISSATPYKAICTGVINMVLVS 61
OY	63 TTRPOLNCTRTSTTPRCPPSSRTENNYICVKCENQPVHFAIGRCP 111
DB	62 TTRPOLNCTRTSTTPRCPPSSRTENNYICVKCENQPVHFAIGRCP 110
RESULT 7	
AAV33321	
ID	AAV33321 standard; Protein: 111 AA.
XX	
AC	AAV33321:
XX	
DT	29-NOV-1999 (first entry)
XX	
DE	Frog lectin protein fragment.
XX	
KM	Cytotoxic; RNase; ribonuclease; pancreatic; antibody; light chain;
KM	heavy chain; cell surface marker; treatment; tumor; viral infection;
KM	parasitic infection; immune dysfunctional cell; autoimmune disease;
KM	contraceptive; cell separation; transplantation; bone marrow ablation;
KM	leukemia cell; T-cell; graft-versus-host disease; bullfrog; lectin.
XX	
OS	Rana catesbeiana.
XX	
PN	US955073-A.
XX	
PD	21-SEP-1999.
XX	
PF	09-JUL-1997; 97US-0891848.
XX	
PR	22-SEP-1993; 93US-0125462.
PR	22-OCT-1991; 91US-0779195.
PR	20-APR-1990; 90US-0510696.
PR	04-FEB-1993; 93US-0014082.
XX	
PA	(USSH ) US DEPT HEALTH & HUMAN SERVICES.
XX	
PI	Rybak SM, Newton DL, Nicholls PJ, Youle RJ;
XX	
DR	WPI; 1999-560488/47.
XX	
PT	Recombinantly fused pancreatic RNase-targeting proteins useful for
PT	treating tumors, infections, immune or autoimmune disorders and as a
PT	contraceptive -
XX	
PS	Example 3; Fig 19; 47pp: English.
XX	
CC	This invention describes a novel nucleic acid construct comprising
CC	sequences encoding functional pancreatic RNase and a second protein
CC	(preferably the light and heavy chains of an antibody) which binds a

CC specific cell surface marker on a target cell and functions as a  
 CC cytotoxic agent. The products can be used for selectively killing cells  
 CC expressing a specific surface marker. They can be used for treating  
 CC tumors or infected cells (e.g. cells infected by viruses (especially  
 CC latent or chronic virus infections, such as human immunodeficiency virus  
 CC (HIV)-1, Epstein-Barr virus, herpes viruses (herpes simplex types 1 and  
 CC II), hepatitis viruses (B, non-A-non-B, and delta), herpes zoster,  
 CC cytomegalovirus) and cells infected with parasites (such as the malaria  
 CC parasite). They can also be used for treating immune dysfunctional cells  
 CC in immune and autoimmune diseases. Additionally, they may be used as  
 CC contrast agents. Finally they can also be used for cell separation in  
 CC vitro by selectively killing unwanted types of cells (e.g. in bone  
 CC marrow) prior to transplantation into a patient undergoing marrow  
 CC ablation by radiation or for killing leukemia cells or T-cells that would  
 CC cause graft-versus-host disease. This sequence represents a bullfrog  
 CC (Rana catesbeiana) lectin used to describe the method of the invention.

XX Sequence 111 AA:

Query Match 96.3%; Score 582.5; DB 20; Length 111;  
 Best Local Similarity 96.4%; Pred. No. 2.5e-58;  
 Matches 107; Conservative 3; Mismatches 0; Indels 1; Gaps 1;

OY 2 QNMAIFQOKHIIITPII-CNFIIDNNIYIVGGCKRVNTFISSATTVAICTGVINLV 60  
 :|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:  
 Db 1 ENMAIFQOKHIIITPIIINCNITMDNNIYIVGGCKRVNTFISSATTVAICTGVINLV 60  
 OY 61 LSTRFQNLCTRTSITPRCPYSSRTETNYICVKNOCNPVHAGIGRCP 111  
 :|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:  
 Db 61 LSTRFQNLCTRTSITPRCPYSSRTETNYICVKNOCNPVHAGIGRCP 111

#### RESULT 8

AAV28869  
 ID AAV28869 standard; Protein: 105 AA.

AC AAV28869;

DT 25-JAN-2000 (first entry)

XX Recombinant Met(-1) RapLRI Met23Leu-(His)6 protein.

XX Recombinant Met(-1) Rana pipiens ribonuclease Met23Leu-(His)6; RapLRI;  
 KW CD22; covalently bound; IL2 antibody; ligand binding moiety; RNase;  
 KW cancerous B cell; Kaposi's sarcoma; human chorionic gonadotropin; hCG;  
 KW signal peptide; recombinant ribonuclease; cytotoxic fusion protein;  
 KW cancer; frog; autoimmune disease.

XX Rana pipiens.  
 OS Synthetic.

XX Key Location/Qualifiers

FT Misc-difference 1 /note- "His)6 histidine tag attached to N-terminal Met"

FT Misc-difference 1 /note- "Met not found in wild type RapLRI"

FT Misc-difference 24 /note- "Wild type Met replaced with Leu"

XX WO9950398-A2.

XX 07-OCT-1999.

XX 26-MAR-1999; 99WO-US06641.

XX 27-MAR-1998; 98US-0079751.

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.

XX Newton DL, Rybak SM;

XX WPI: 1999-610847/52.

XX N-PSDB: AAZ08127

XX New recombinant ribonucleases, used for killing target cells, e.g. for  
 PT treating cancers, viral infections or autoimmune diseases  
 XX Claim 4; Page 59; 71pp; English.

XX The present sequence is a recombinant Rana pipiens ribonuclease protein  
 CC (RapLRI) with Met at position 1 attached to (His)6 tag and Met24Leu.  
 CC Carboxy terminal end of recombinant RapLRI has a covalently bound ligand  
 CC binding moiety, which can be a IL2 antibody directed against CD22 on  
 CC cancerous B cells or human chorionic gonadotropin (hCG) effective  
 CC against Kaposi's sarcoma cells. Recombinant ribonucleases can be  
 CC expressed in bacteria without an N-terminal methionine due to the  
 CC presence of a signal peptide that is cleaved by bacteria. The soluble  
 CC expression of ribonuclease allows the proteins to be fused in-frame with  
 CC ligand binding moieties to form cytotoxic fusion proteins. They can be  
 CC used for treatment of cancer and autoimmune diseases.

XX Sequence 105 AA:

Query Match 47.0%; Score 284.5; DB 20; Length 105;  
 Best Local Similarity 50.0%; Pred. No. 1.3e-24;  
 Matches 56; Conservative 15; Mismatches 32; Indels 9; Gaps 4;

OY 1 QNMAIFQOKHIIITPII-CNFIIDNNIYIVGGCKRVNTFISSATTVAICTGVINLV 58  
 :|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:  
 Db 1 QMDLIFQOKHIIITPIIINCNITMDNNIYIVGGCKRVNTFISSATTVAICTGVINLV 56  
 OY 59 NVLSTRFQNLCTRTSITPRCPYSSRTETNYICVKNOCNPVHAGIGRCP 110  
 :|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:|||||:  
 Db 57 NVLSTRFQNLCTRTSITPRCPYSSRTETNYICVKNOCNPVHAGIGRCP 105

#### RESULT 9

AAV28867  
 ID AAV28867 standard; Protein: 105 AA.

AC AAV28867;

DT 25-JAN-2000 (first entry)

XX Recombinant Met(-1) RapLRI.

XX Recombinant Met(-1) Rana pipiens ribonuclease; RapLRI; CD22; RNase;  
 KW covalently bound; IL2 antibody; ligand binding moiety; cancerous B cell;  
 KW Kaposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;  
 KW recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;  
 KW autoimmune disease.

XX Rana pipiens.  
 OS Synthetic.

XX Key Location/Qualifiers

FT Misc-difference 1 /note- "Met not found in wild type RapLRI"

XX WO9950398-A2.

XX 07-OCT-1999.

XX 26-MAR-1999; 99WO-US06641.

XX 27-MAR-1998; 98US-0079751.

XX (USSH ) US DEPT HEALTH & HUMAN SERVICES.

XX Newton DL, Rybak SM;

XX WPI: 1999-610847/52.

XX N-PSDB: AAZ08126.

XX New recombinant ribonucleases, used for killing target cells, e.g. for  
 PT treating cancers, viral infections or autoimmune diseases

XX Claim 34; Page 57; 71pp; English.  
 PS  
 CC The present sequence is a recombinant Rana pipiens ribonuclease (RaplR1)  
 CC protein with Met at position 1. Carboxy terminal end of recombinant  
 CC RaplR1 has a covalently bound ligand binding moiety, which can be a IL2  
 CC antibody directed against CD22 on cancerous B cells or human chorionic  
 CC gonadotrophin (hCG) effective against Kaposi's sarcoma cells. Recombinant  
 CC ribonucleases can be expressed in bacteria without an N-terminal  
 CC methionine due to the presence of a signal peptide that is cleaved by  
 CC bacteria. The soluble expression of ribonuclease allows the proteins to  
 CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
 CC proteins. They can be used for treatment of cancer and autoimmune  
 CC diseases.  
 CC  
 SO Sequence 105 AA;  
 Query Match 46.7%; Score 282.5; DB 20; Length 105;  
 Best Local Similarity 49.1%; Pred. No. 2.5e-24;  
 Matches 55; Conservative 16; Mismatches 32; Indels 9; Gaps 4;  
 QY 1 MONWATFOOKHIINT-PIICNTILDNNIYIVGGCKRVNFISSATTVAICIGVI-NL 58  
 DB 1 MODWLFQKKHNTFVDVDCNNIMSTNLF---HCKDKNFTYSRPEPKAICKGIASK 56  
 QY 59 NVLSTFRQLNTCTRTSTTPRCPYSSRTETNYICVKNQYPAVFAGIGRC 110  
 DB 57 NVLTSEFYLSDC---NVTSRPKYKLRKSTNFCVTCENAPVHFVGVGHC 105  
 RESULT 10  
 AAW06544  
 ID AAW06544 standard; protein: 104 AA.  
 AC AAW06544;  
 XX  
 DT 22-AUG-1997 (first entry)  
 DE Antitumour protein from Rana pipiens oocytes.  
 XX  
 KM Tumour; chemotherapy; radiotherapy; frog.  
 XX  
 OS Rana pipiens.  
 XX  
 PN WO9639428-A1.  
 PD 12-DEC-1996.  
 XX  
 PF 03-JUN-1996; 96WO-US08304.  
 XX  
 PR 06-JUN-1995; 95US-0467955.  
 XX  
 PA (ALFA-) ALFACELL CORP.  
 XX  
 PI Ardelt WJ;  
 XX  
 DR WPI; 1997-043063/04.  
 XX  
 PT Antitumour proteins from Rana pipiens oocyte(s) - have fewer  
 PS disadvantages than chemotherapy, surgery and radiotherapy  
 XX  
 CC Claim 8; Page 28; 45pp; English.  
 XX  
 CC The present sequence is a specifically claimed example of an  
 CC antitumour protein from the generic protein in AAW18224, with the  
 CC molecular weight 12000. This is one of two preferred proteins (the  
 CC other in AAW06543) that have been isolated from Rana pipiens oocytes.  
 CC Both proteins have a blocked amino terminal group and are essentially  
 CC free of carbohydrates. The proteins are used to treat tumours. Use of  
 CC the peptides has fewer disadvantages than chemotherapy, radiotherapy  
 CC and surgery in the treatment of tumours.  
 CC  
 SO Sequence 104 AA;

Query Match 46.4%; Score 280.5; DB 18; Length 104;  
 Best Local Similarity 48.6%; Pred. No. 4.2e-24;  
 Matches 54; Conservative 17; Mismatches 31; Indels 9; Gaps 4;  
 QY 2 QMWATFOOKHIINT-PIICNTILDNNIYIVGGCKRVNFISSATTVAICIGVI-NLN 59  
 DB 1 EDWLTFOKKHNTFVDVDCNNIMSTNLF---HCKDKNFTYSRPEPKAICKGIASKN 56  
 QY 60 VLSTFRQLNTCTRTSTTPRCPYSSRTETNYICVKNQYPAVFAGIGRC 110  
 DB 57 VLTSEFYLSDC---NVTSRPKYKLRKSTNFCVTCENAPVHFVGVGHC 104  
 RESULT 11  
 AAY28866  
 ID AAY28866 standard; protein: 104 AA.  
 AC AAY28866;  
 XX  
 DT 25-JAN-2000 (first entry)  
 DE Recombinant RaplR1 Met23leu amino acid sequence.  
 XX  
 KM Recombinant Rana pipiens ribonuclease; RaplR1 Met23leu; covalently bound;  
 KM IL2 antibody; ligand binding moiety; CD22; cancerous B cell; RNase;  
 KM Kaposi's sarcoma; human chorionic gonadotrophin; hCG; signal peptide;  
 KM recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;  
 KM autoimmune disease.  
 XX  
 OS Rana pipiens.  
 OS Synthetic.  
 XX  
 FH Key location/qualifiers  
 FT Misc-difference 23 /note- "wild type Met replaced with Leu"  
 FT  
 XX  
 PN WO950398-A2.  
 PD 07-OCT-1999.  
 XX  
 PF 26-MAR-1999; 99WO-US06641.  
 XX  
 PR 27-MAR-1998; 98US-0079751.  
 XX  
 PA (USSH ) US DEPT HEALTH & HUMAN SERVICES.  
 XX  
 PI Newton DL, Rybak SM;  
 XX  
 DR WPI; 1999-610847/52.  
 DR N-PSDB; AA208125.  
 XX  
 PT New recombinant ribonucleases, used for killing target cells, e.g. for  
 PS treating cancers, viral infections or autoimmune diseases  
 XX  
 CC Claim 34; Page 56; 71pp; English.  
 XX  
 CC The present sequence is a recombinant Rana pipiens ribonuclease (RaplR1)  
 CC protein with Met23leu. Carboxy terminal end of recombinant RaplR1 has a  
 CC covalently bound ligand binding moiety, which can be a IL2 antibody  
 CC directed against CD22 on cancerous B cells or human chorionic  
 CC gonadotrophin (hCG) effective against Kaposi's sarcoma cells. Recombinant  
 CC ribonucleases can be expressed in bacteria without an N-terminal  
 CC methionine due to the presence of a signal peptide that is cleaved by  
 CC bacteria. The soluble expression of ribonuclease allows the proteins to  
 CC be fused in-frame with ligand binding moieties to form cytotoxic fusion  
 CC proteins. They can be used for treatment of cancer and autoimmune  
 CC diseases.  
 CC  
 SO Sequence 104 AA;  
 Query Match 46.2%; Score 279.5; DB 20; Length 104;  
 Best Local Similarity 49.5%; Pred. No. 5.5e-24;

OY	1	MOMNATFOQKHIIINT-PIICENTILDDNIIVYGCGCKRVNFETISSATVAKITGYI-NL	58
Db	1	MODULTFQKHHINTKIKVDCCDINIMSTLF---HCKKNFETISRPBPAVKIGKIASK	56
OY	59	NVLSTFTFOQNTCTRTSITRPPCPYSRTETNYICVACENQOYVVHAGIGRC	110
Db	57	NVLTTSEFYSDC---NVTSRPPCKIKLKSKTKNFCVTGCENQAOPVHFEGVASC	105

RESULT 13  
AAZ28865

ID	AAV28865	standard; protein; 104	AA
XX			

XX

XX

XX  
Page 1 of 1

KW human chorionic gonadotropin: hCG: recombinant ribonuclease: RNase: ligand binding moiety; CD22; cancerous B cell; Kaposi's sarcoma; Krog;

signal peptide; cytolotoxic fusion protein; cancer; autoimmune disease.

XX

XX

**XX**

**XX**

XX 27-MAR-1998. 08115-0079751  
PR

PA (USSH ) US DEPT HEALTH & HUMAN SERVICES  
AA

PI Newton DL, Rybak SM;

DR WPI; 1999-610847/52.

100

treating cancers, viral infections

ps Claim 1; Page 55; 1pp; English.  
XX

CC The present sequence is RALPIRPI has a covalently bound  
CC protein. Carboxy terminal end of RALPIRPI has a covalently bound

CD22 on cancerous B cells or human chorionic gonadotrophin (hCG)

CC be expressed in bacteria without an N-terminal methionine due to the

CC expression of ribonuclease allows the proteins to be fused in-frame with

CC used for treatment of cancer and autoimmune diseases.

sequence 104 AA;

Query Machine	Score	DB Size	Length
Best Local Similarity	48.68;	Pred. No. 9.3e-24;	

[illegible]

60 110

Db 57 VI.TTSEFYI.SDC---NV.TSPBCKYKI.KKSTNTCEVTCE.NADPVHEVGVGHC 70A

RESULT	14
ID	AAIY28671 standard; Protein: 105 AA.
AC	AAIY28671:
XX	
XX	
DT	25-JAN-2000 (first entry)
XX	
DE	Recombinant Met(-1) RapLRI GlnSer amino acid sequence.
KW	Recombinant Met(-1) Rana pipiens ribonuclease GlnSer; RAPLRI; CD22;
KW	covalently bound; IL2 antibody; ligand binding moiety; cancerous B cell;
KM	Raposi's sarcoma; human chorionic gonadotropin; hCG; signal peptide;
KV	recombinant ribonuclease; cytotoxic fusion protein; cancer; frog;
KX	autoimmune disease; RNase.
OS	Rana pipiens.
OS	Synthetic.
FH	Key
FT	Misc-difference 1 Location/Qualifiers
FT	/note= "Met not found in wild type RapLRI"
FT	Misc-difference 2 /note= "wild type Gln replaced with Ser"
ET	
PX	WO9950398-A2.
PN	
PD	07-OCT-1999.
XX	
PP	26-MAR-1999; 99WO-US06641.
XX	
PR	27-MAR-1998; 98US-0079751.
PA	(USSH ) US DEPT HEALTH & HUMAN SERVICES.
PI	Newton DL, Rybak SM;
XX	
DR	WPI: 1999-610847/52.
N-PSDB:	AAZ08129.
XX	
PT	New recombinant ribonucleases, used for killing target cells, e.g. for
PT	treating cancers, viral infections or autoimmune diseases -
XX	
PS	Claim 34; Page 61; 71pp; English.
XX	
CC	The present sequence is a recombinant Rana pipiens ribonuclease (RAPLRI)
CC	protein with Met at position 1 and Gln2Ser. Carboxy terminal end of
CC	recombinant RAPLRI has a covalently bound ligand binding moiety, which
CC	can be a IL2 antibody directed against CD22 on cancerous B cells or human
CC	chorionic gonadotropin (hCG) effective against Kaposi's sarcoma cells.
CC	Recombinant ribonucleases can be expressed in bacteria without an N-
CC	terminal methionine due to the presence of a signal peptide that is
CC	cleaved by bacteria. The soluble expression of ribonuclease allows the
CC	proteins to be fused in-frame with ligand binding moieties to form
CC	cytotoxic fusion proteins. They can be used for treatment of cancer and
CC	autoimmune diseases.
SO	Sequence 105 AA:
OY	Query Match 45.9%; Score 277.5; DB 20; Length 105;
Best Local Similarity:	48.2%; Pred.No. 9.4e-24;
Matches 54; Conservative 16; Mismatches 33; Indels 9; Gaps 4	
Db .	1 MONNATPQOKKIINT-PICHTIIDNNIIYVGCGCKRNVTFISATTVAICTGVT-NL 58
I MSDMTFCKKHLTNRDVCNNINSTNF---HCKDKNTFIYSRPEPVKAICKGIASK 56	
OY	59 NVLTSTRQLTMTCRTSITPPCPSSRTETNYIVCKENOXYPVFAGIGRC 110
Db	57 NVLTSTSEYLDSC---NVTSRPKCKKKLKSKSTNTFCVTECNOAPAFHFVGVC 105
RESULT	15

[illegible]



Wed Jun 25 15:53:46 2003

Search completed: June 25, 2003, 14:48:41  
job time : 34.3 secs

us-09-622-613b-21.rag

